Connecting via Winsock to STN

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Welcome to STN International! Enter x:X
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LOGINID:ssspta1604dxj

PASSWORD:

TERMINAL (ENTER 1, 2, 3, OR ?):2

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                                                    * * * * * * * * * *
                     Welcome to STN International
                 Web Page for STN Seminar Schedule - N. America
NEWS 1
 NEWS
      2 AUG 15 CAOLD to be discontinued on December 31, 2008
 NEWS
      3 OCT 07 EPFULL enhanced with full implementation of EPC2000
NEWS 4 OCT 07 Multiple databases enhanced for more flexible patent
                 number searching
NEWS 5 OCT 22 Current-awareness alert (SDI) setup and editing
                 enhanced
NEWS 6 OCT 22 WPIDS, WPINDEX, and WPIX enhanced with Canadian PCT
                 Applications
 NEWS
         OCT 24 CHEMLIST enhanced with intermediate list of
                 pre-registered REACH substances
 NEWS 8 NOV 21 CAS patent coverage to include exemplified prophetic
                 substances identified in English-, French-, German-,
                 and Japanese-language basic patents from 2004-present
NEWS 9 NOV 26 MARPAT enhanced with FSORT command
NEWS 10 NOV 26 MEDLINE year-end processing temporarily halts
                 availability of new fully-indexed citations
 NEWS 11 NOV 26
                 CHEMSAFE now available on STN Easy
 NEWS 12 NOV 26 Two new SET commands increase convenience of STN
                 searching
NEWS 13 DEC 01 ChemPort single article sales feature unavailable
NEWS 14 DEC 12 GBFULL now offers single source for full-text
                 coverage of complete UK patent families
NEWS 15 DEC 17 Fifty-one pharmaceutical ingredients added to PS
NEWS EXPRESS JUNE 27 08 CURRENT WINDOWS VERSION IS V8.3,
             AND CURRENT DISCOVER FILE IS DATED 23 JUNE 2008.
NEWS HOURS
              STN Operating Hours Plus Help Desk Availability
NEWS LOGIN
              Welcome Banner and News Items
NEWS IPC8
              For general information regarding STN implementation of IPC 8
Enter NEWS followed by the item number or name to see news on that
```

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FILE 'HOME' ENTERED AT 16:34:41 ON 30 DEC 2008

specific topic.

```
=> file reg
COST IN U.S. DOLLARS
                                                 SINCE FILE
                                                                 TOTAL
                                                             SESSION
                                                     ENTRY
FULL ESTIMATED COST
                                                       0.21
                                                                  0.21
FILE 'REGISTRY' ENTERED AT 16:35:03 ON 30 DEC 2008
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COPYRIGHT (C) 2008 American Chemical Society (ACS)
Property values tagged with IC are from the ZIC/VINITI data file
provided by InfoChem.
STRUCTURE FILE UPDATES:
                          29 DEC 2008 HIGHEST RN 1091682-77-7
DICTIONARY FILE UPDATES: 29 DEC 2008 HIGHEST RN 1091682-77-7
New CAS Information Use Policies, enter HELP USAGETERMS for details.
TSCA INFORMATION NOW CURRENT THROUGH July 5, 2008.
  Please note that search-term pricing does apply when
 conducting SmartSELECT searches.
REGISTRY includes numerically searchable data for experimental and
predicted properties as well as tags indicating availability of
experimental property data in the original document. For information
on property searching in REGISTRY, refer to:
http://www.cas.org/support/stngen/stndoc/properties.html
=> s lemuteporfin/cn
           1 LEMUTEPORFIN/CN
L1
=> d 11
    ANSWER 1 OF 1 REGISTRY COPYRIGHT 2008 ACS on STN
    215808-49-4 REGISTRY
RN
     Entered STN: 17 Dec 1998
F.D
     23H, 25H-Benzo[b] porphine-9, 13-dipropanoic acid,
     18-ethenyl-4,4a-dihydro-3,4-bis(methoxycarbonyl)-4a,8,14,19-tetramethyl-,
     9,13-bis(2-hydroxyethyl) ester (CA INDEX NAME)
OTHER CA INDEX NAMES:
     23H, 25H-Benzo[b]porphine-9, 13-dipropanoic acid,
     18-ethenyl-4, 4a-dihydro-3, 4-bis (methoxycarbonyl) -4a, 8, 14, 19-tetramethyl-,
     bis(2-hydroxyethyl) ester (9CI)
OTHER NAMES:
CN
    A-EA 6
CN
     EA 6
CN
     Lemuteporfin
CN
     OLT 0074
MF
     C44 H48 N4 O10
SR
```

STN Files: ADISINSIGHT, BIOSIS, CA, CAPLUS, IMSDRUGNEWS, IMSRESEARCH,

PROUSDDR, TOXCENTER, USAN, USPAT2, USPATFULL

L.C.

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Ме
                                             OM∈
            H2C== CH-
                          Ν
                              HN
                                            OMe
                                     Ме
                                         0
                           ΝН
                                N-
                                       Me
                                             0
                                       - cH<sub>2</sub>- c- о- сH<sub>2</sub>- сH<sub>2</sub>- он
но-сн2-сн2-о-с-сн2-
                       CH<sub>2</sub>
                                   CH2
**PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT**
               28 REFERENCES IN FILE CA (1907 TO DATE)
               1 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
               28 REFERENCES IN FILE CAPLUS (1907 TO DATE)
=> s vertporfin/cn
             0 VERTPORFIN/CN
T.2
=> s verteporfin/cn
             1 VERTEPORFIN/CN
L3
=> d 13
     ANSWER 1 OF 1 REGISTRY COPYRIGHT 2008 ACS on STN
L3
     129497-78-5 REGISTRY
RN
     Entered STN: 21 Sep 1990
ED
     24H, 26H-Benzo[b]porphine-9, 13-dipropanoic acid,
     18-ethenyl-4,4a-dihydro-3,4-bis(methoxycarbonyl)-4a,8,14,19-tetramethyl-,
     monomethyl ester, (4R, 4aS)-rel- (CA INDEX NAME)
OTHER CA INDEX NAMES:
     23H, 25H-Benzo[b] porphine-9, 13-dipropanoic acid,
     18-ethenyl-4,4a-dihydro-3,4-bis(methoxycarbonyl)-4a,8,14,19-tetramethyl-,
     monomethyl ester, trans-
OTHER NAMES:
CN
     BPD-MA
CN
     CL 318952
CN
     Verteporfin
CN
     Visudyne
FS
     STEREOSEARCH
DR
     121987-00-6, 129162-83-0, 136415-38-8
MF
     C41 H42 N4 O8
CI
     IDS
SR
                  ADISINSIGHT, ADISNEWS, AGRICOLA, BIOSIS, CA, CAPLUS, CBNB,
LC
       CHEMCATS, CIN, DDFU, DRUGU, EMBASE, IMSDRUGNEWS, IMSPATENTS, IMSPRODUCT,
       IMSRESEARCH, IPA, MEDLINE, MRCK*, PATDPASPC, PHAR, PROMT, PROUSDDR, PS,
       RTECS*, TOXCENTER, USAN, USPAT2, USPATFULL
         (*File contains numerically searchable property data)
     Other Sources:
     CM
          1
     CRN 121310-58-5
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10588419

CMF C40 H40 N4 O8

Relative stereochemistry.

CM 2

CRN 67-56-1 CMF C H4 O

 $_{
m H3C}-_{
m OH}$

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

524 REFERENCES IN FILE CA (1907 TO DATE)

25 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

525 REFERENCES IN FILE CAPLUS (1907 TO DATE)

=> file medicine
FILE 'DRUGMONOG' ACCESS NOT AUTHORIZED
COST IN U.S. DOLLARS

COST IN U.S. DOLLARS
SINCE FILE TOTAL
ENTRY SESSION
FULL ESTIMATED COST
19.91
20.12

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FILE 'USGENE' ENTERED AT 16:35:55 ON 30 DEC 2008 COPYRIGHT (C) 2008 SEQUENCEBASE CORP

FILE 'USPATFULL' ENTERED AT 16:35:55 ON 30 DEC 2008
CA INDEXING COPYRIGHT (C) 2008 AMERICAN CHEMICAL SOCIETY (ACS)

FILE 'USPATOLD' ENTERED AT 16:35:55 ON 30 DEC 2008
CA INDEXING COPYRIGHT (C) 2008 AMERICAN CHEMICAL SOCIETY (ACS)

FILE 'USPAT2' ENTERED AT 16:35:55 ON 30 DEC 2008
CA INDEXING COPYRIGHT (C) 2008 AMERICAN CHEMICAL SOCIETY (ACS)

- => s 11 or lemuteporfin or qlt 0074
- 'CN' IS NOT A VALID FIELD CODE
- 'CN' IS NOT A VALID FIELD CODE 'CN' IS NOT A VALID FIELD CODE
- 'CN' IS NOT A VALID FIELD CODE
- 'CN' IS NOT A VALID FIELD CODE
- L4 145 L1 OR LEMUTEPORFIN OR QLT 0074
- => s 13 or verteporfin or visudyne or CL 318952
- 'CN' IS NOT A VALID FIELD CODE
- 'CN' IS NOT A VALID FIELD CODE
- 'CN' IS NOT A VALID FIELD CODE

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'CN' IS NOT A VALID FIELD CODE
          8048 L3 OR VERTEPORFIN OR VISUDYNE OR CL 318952
=> s lipophilic
       204158 LIPOPHILIC
=> s acne or seborrheic dermatitis or hyperactive sebaceous gland or sebaceous gland
hyperplasia or seborrhea
        208769 ACNE OR SEBORRHEIC DERMATITIS OR HYPERACTIVE SEBACEOUS GLAND OR
               SEBACEOUS GLAND HYPERPLASIA OR SEBORRHEA
=> s 14 or 15
         8121 L4 OR L5
L8
=> s 18 and 17
L9
          209 L8 AND L7
=> s photodynamic therapy or PDT
        80855 PHOTODYNAMIC THERAPY OR PDT
=> s 19 and 110
        62 L9 AND L10
T.11
=> s hydrophobic
      898093 HYDROPHOBIC
=> s 15 or 112
L13 905604 L5 OR L12
=> s 111 and 113
           60 L11 AND L13
L14
=> dup rem
ENTER L# LIST OR (END):114
DUPLICATE IS NOT AVAILABLE IN 'ADISINSIGHT, ADISNEWS, DGENE, DRUGMONOG2,
IMSPRODUCT, KOSMET, NUTRACEUT, PCTGEN, PHARMAML, USGENE'.
ANSWERS FROM THESE FILES WILL BE CONSIDERED UNIQUE
PROCESSING COMPLETED FOR L14
L15
             55 DUP REM L14 (5 DUPLICATES REMOVED)
=> s 115 and pd<2004
   5 FILES SEARCHED...
'2004' NOT A VALID FIELD CODE
'2004' NOT A VALID FIELD CODE
'2004' NOT A VALID FIELD CODE
 14 FILES SEARCHED...
 16 FILES SEARCHED...
'2004' NOT A VALID FIELD CODE
 22 FILES SEARCHED...
'2004' NOT A VALID FIELD CODE
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27 FILES SEARCHED...

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'2004' NOT A VALID FIELD CODE
 31 FILES SEARCHED...
            9 L15 AND PD<2004
=> d 116 1-9 ibib, kwic
L16 ANSWER 1 OF 9 CAPLUS COPYRIGHT 2008 ACS on STN
ACCESSION NUMBER:
                         2003:836896 CAPLUS
DOCUMENT NUMBER:
                          139:288313
                          High fluence rate activation of photosensitizers for
TITLE:
                          dermatological applications
INVENTOR(S):
                          Geronemus, Roy G.; Alexiades-Armenakas, Macrene;
                          McMillan, Kathleen
PATENT ASSIGNEE(S):
                          Candela Corporation, USA
SOURCE:
                          PCT Int. Appl., 30 pp.
                          CODEN: PIXXD2
DOCUMENT TYPE:
                           Patent
LANGUAGE:
                           English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:
     PATENT NO.
                     KIND DATE APPLICATION NO. DATE
                                  -----
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                          ____
                                               _____
                                                                        _____
     WO 2003086460 A2 20031023
WO 2003086460 A3 20031231
                          A2 20031023 WO 2003-US10418
                                                                     20030404 <--
         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
              CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
              GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
              LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM,
              PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT,
              TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
                          A1 20031027
                                            AU 2003-230808 20030404 <--
     AU 2003230808
                                               US 2002-370253P P 20020405
WO 2003-US10418 W 20030404
PRIORITY APPLN. INFO.:
REFERENCE COUNT:
                                 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS
                           2.
                                 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
     WO 2003086460 A2 20031023
                        KIND DATE
                                            APPLICATION NO.
                                                                       -----
     _____
                          ----
                                               -----
                                             WO 2003-US10418
     WO 2003086460 A2 20031023 WO 2003086460 A3 20031231
                                                                       20030404 <--
PΤ
                          A3 20031231
     WO 2003086460
         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
              CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM,
              PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT,
              TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
              KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES,
              FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR,
              BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
     AU 2003230808 A1 20031027 AU 2003-230808
     . . does not cause clin. signification side effects such as purpura
     of the treated skin. Examples of photosensitizers used are
     \delta-aminolevulinate, <u>verteporfin</u> and hypericin for treatment
```

```
of actinic keratosis, acne, basal cell carcinoma, photoaged
     skin, and rosacea. The treatment is also suitable for hair removal.
     skin disease photodynamic treatment; aging skin photodynamic
ST
     therapy; hair removal photodynamic treatment
ΙT
     Acne
     Antitumor agents
     Eczema
     Human
     Hyperthermia (therapeutic)
      Photodynamic therapy
     Photosensitizers, pharmaceutical
     Psoriasis
     Radio wave
     Skin, disease
     Skin, neoplasm
     Wart
        (high fluence rate activation of photosensitizers for dermatol.
        applications)
ΙT
     Acne
        (vulgaris; high fluence rate activation of photosensitizers for
        dermatol. applications)
                                  92-83-1D, Xanthene, derivs.
TT
     81-54-9D, Purpurin, derivs.
     \delta-Aminolevulinic acid 548-04-9, Hypericin 2683-78-5D,
     Bacteriochlorin, derivs. 20238-92-0, N-Acetyl \delta-aminolevulinic
          33320-16-0 73442-88-3 129497-78-5, Verteporfin
     140898-98-2
                  149837-93-4D, Bacteriopurpurin, derivs. 186410-03-7
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (high fluence rate activation of photosensitizers for dermatol.
        applications)
L16 ANSWER 2 OF 9 COPYRIGHT 2008 Gale Group on STN
ACCESSION NUMBER:
                    2003:225291 NLDB
                    OTHER NEWS TO NOTE.
TITLE:
SOURCE:
                    BIOWORLD Today, (18 Nov 2003) .
                    Medical Economics/Thomson Healthcare
PUBLISHER:
DOCUMENT TYPE:
                    Newsletter
LANGUAGE:
                    English
WORD COUNT:
                    3388
     BIOWORLD Today, (18 \text{ Nov } 2003) .
        Micrologix . . from its Phase IIb study of MBI 594AN, a topical
     drug candidate under development as a first-in-class prescription
     treatment for acne. The Phase II study was designed to evaluate
     acne lesion count reductions at various time points (three, six,
     nine and 12 weeks), comparing MBI 594AN (1.25 percent and 2.5. .
        Miravant . . . cause of blindness in adults more than 50 years old.
     Safety data showed that the proposed clinical dose of SnET2-PhotoPoint
     photodynamic therapy was well tolerated and demonstrated
a favorable profile in the study population.
        QLT . . . American Academy of Ophthalmology meeting in Anaheim,
     Calif., showed that Macugen does not appear to provide an improvement over
     QLT's Visudyne Therapy for patients with choroidal
     neovascularization due to age-related macular degeneration. QLT noted that
     although complete data were not presented, the anti-VEGF aptamer data
     appear no better than Visudyne's original TAP data in all lesion
     types. Macugen (pegaptanib sodium) is under development by Eyetech
     Pharmaceuticals Inc., of New York, . . .
```

L16 ANSWER 3 OF 9 COPYRIGHT 2008 Gale Group on STN

Medical Review Criteria Guidelines for Managing Care.

M2 Presswire, $(\underline{19} \text{ Feb } \underline{2003})$.

SOURCE: M2 Presswire, (19 Feb M2 Communications Ltd.
DOCUMENT TYPE: Newsletter
LANGUAGE: English
WORD COUNT: 3984
SO M2 Presswire

M2 Presswire, (19 Feb 2003) .

Dermatology: Dermatology referral management; Acne vulgaris; Actinic keratosis; Alopecia areata; Atopic dermatitis; Basal cell carcinoma; Biopsy/excision of benign skin and subcutaneous lesions, Cysts involving the. . .

Laser . . . Micrographic Surgery; Pediculosis (`lice`); Photochemotherapy for the treatment of scleroderma, extracorporeal; Psoriasis; PUVA Therapy; Rosacea; Scabies; Sclerotherapy for varicose veins, Seborrheic dermatitis/`Dandruff`; Seborrheic keratosis; Squamous cell carcinoma; Tattoos; Verruca Vulgaris/ Warts; Vitiliqo;

Age-related Macular Degeneration (AMD); Macular Degeneration, Radiation Treatment; Macular Fovea Translocation for AMD; Ocular Photodynamic Therapy (OPT) - Visudyne (Verteporfin) Therapy for Age-related Macular Degeneration; Opthalmoscopy, extended with retinal drawing Orbitotomy; Photocoagulation;

Pterygium; Ptosis; Punctal dilation/snipping; Punctal Plugs and Punctoplasty. . . Thermal Therapy; Age- Related Macular Degeneration; Macular Degeneration, Laser Therapy; ar Degeneration, Radiation Treatment; Macular Fovea Translocation for AMD; Ocular Photodynamic

Therapy (OPT) - Visudyne (Verteporfin) Therapy for Age-related Macular Degeneration; Viscocanalostomy; Visual Field

Testing; Visual training (Orthoptics); Visual Rehabilitation Program; Vitrectomy; Referral criteria summary sheet/grid.

L16 ANSWER 4 OF 9 USPATFULL on STN

ACCESSION NUMBER: 2003:300363 USPATFULL

TITLE: Keptin-a novel keratinocyte-specific proteinase

inhibitor

INVENTOR(S): Ariizumi, Kiyoshi, Plano, TX, UNITED STATES Cruz, Ponciano D., Dallas, TX, UNITED STATES

Board of Regents, The University of Texas System (U.S. PATENT ASSIGNEE(S):

corporation)

NUMBER KIND DATE US 20030211587 A1 20031113 US 2002-141530 A1 20020507 (10) PATENT INFORMATION: <--APPLICATION INFO.: DOCUMENT TYPE: Utility

APPLICATION FILE SEGMENT:

LEGAL REPRESENTATIVE: Steven L. Highlander, FULBRIGHT & JAWORSKI L.L.P., Suite 2400, 600 Congress Avenue, Austin, TX, 78701

NUMBER OF CLAIMS: 47 EXEMPLARY CLAIM:

3 Drawing Page(s) NUMBER OF DRAWINGS:

LINE COUNT: 1770

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

DETD . . . dermatitis, cutaneous basal cell carcinoma, cutaneous planocellular carcinoma, wart, lameliar ichthyosis, epidemolytic keratosis, solar induced precancerous keratosis, benign keratosis, ache, seborrheic dermatitis, keloids, pityriasis rubra pilaris ("PRP"), dermatomyositis, and angiogenesis-related skin disorders.

. . . weight protein of 12.5 kDa containing no cysteine residues DETD which suggests the formation of a three dimensional structure by leucine-based hydrophobic interactions.

. . amino acids: serine (+0.3), asparagine (+0.2), glutamine DETD (+0.2), and threonine (-0.4), sulfur containing amino acids: cysteine (-1.0) and methionine (-1.3); hydrophobic, nonaromatic amino acids: valine (-1.5), leucine $\overline{(-1.8)}$, isoleucine (-1.8), proline (-0.5 ± 1) , alanine (-0.5), and glycine (0); hydrophobic, aromatic amino acids: tryptophan (-3.4), phenylalanine (-2.5), and tyrosine (-2.3).

DETD . . . dermatitis, cutaneous basal cell carcinoma, cutaneous planocellular carcinoma, wart, lameliar ichthyosis, epidemolytic keratosis, solar induced precancerous keratosis, benign keratosis, ache, seborrheic dermatitis, keloids, pityriasis rubra pilaris ("PRP"), dermatomyositis, angiogenesis-related skin disorders, erysipleas, and erythroderma.

[0113] 5. Phototherapy (UV Irradiation) and Photodynamic DETD Therapy

[0115] Photodynamic therapy (also known as " DETD PDT") involves the administration of a drug followed by light exposure. In PDT, drugs known as porphyrins are administered intravenously into the body to sensitize diseased tissue to visible light. Forms of porphyrin are well known, they include hematoporphyrin derivative (HPD) and porfimer sodium (Photofrin®) and BPD verteporfin.

. . invention, it is contemplated that a nucleic acid segment DETD encoding a keptin may be used in combination with photochemotherapy and photodynamic therapy.

L16 ANSWER 5 OF 9 USPATFULL on STN

2003:153401 USPATFULL ACCESSION NUMBER:

TITLE: Metallotetrapyrrolic photosensitizing agents for use in

photodynamic therapy
Robinson, Byron C., Santa Barbara, CA, UNITED STATES INVENTOR(S):

Leitch, Ian M., Goleta, CA, UNITED STATES Greene, Stephanie, Goleta, CA, UNITED STATES

Rychnovsky, Steve, Santa Barbara, CA, UNITED STATES

NUMBER KIND DATE ______ PATENT INFORMATION: US 20030105069 A1 20030605 <--US 6827926 B2 20041207 US 2002-159005 A1 20020531 (10) APPLICATION INFO.:

> NUMBER DATE ______

US 2001-295345P 20010531 (60) PRIORITY INFORMATION:

DOCUMENT TYPE: Utilitv FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: Finnegan, Henderson, Farabow,, Garrett & Dunner,

L.L.P., 1300 I Street, N.W., Washington, DC, 20005-3315

NUMBER OF CLAIMS: 78 EXEMPLARY CLAIM: 1 7007 LINE COUNT:

- CAS INDEXING IS AVAILABLE FOR THIS PATENT.
- TI Metallotetrapyrrolic photosensitizing agents for use in photodynamic therapy
- SUMM [0002] This invention relates to metallotetrapyrrolic compounds having phototherapeutic properties utilizable in photodynamic therapy for photodetection and phototherapy of target tissues.
- SUMM [0004] Photodynamic therapy ("PDT") is a new modality for the treatment of malignancies, diseased tissue, hyperproliferating tissues, normal tissues or pathogens. PDT involves a localized or systemic administration of a photosensitizing compound followed by exposure of target tissue to photoactivating light. The. . .
- SUMM [0006] An emerging clinical role for <u>photodynamic</u> therapy is in the treatment of proliferative cardiovascular diseases such as atherosclerosis, restenosis and vein graft disease. Atherosclerosis is a disease. . .
- SUMM [0014] Recently, vascular <u>photodynamic</u> therapy has shown promise for the prevention of injury-induced neointimal hyperplasia in animal studies and has entered phase I/II clinical trials. . .
- SUMM . . . cardiovascular field, mostly in preclinical animal models. Such photosensitizers include Photofrin, 5-amino-levulinic acid (protoporphyrin IX precursor), tin ethyl etiopurpurin (SnET2),

 Visudyne® (Benzoporphyrin derivative), Antrin®,

 Optrin® (Lutetium texaphyrin), mono-aspartyl chlorin e6 (MACE), and pheophorbide PH1126. All of these synthetic compounds were designed. .
- SUMM [0016] The excitation light source for \underline{PDT} (usually diode lasers or dye lasers) has historically \overline{been} matched to the far-red absorption bandwidth of the photosensitizer to maximize. . .
- SUMM [0017] Enthusiasm for photoangioplasty (\underline{PDT} of vascular de novo atherosclerotic, restenotic lesions and vein graft intimal hyperplasia) is fueled by more effective second-generation photosensitizers that. . .
- SUMM . . . than 600 nm in the cardiovascular field. This may have been true several years ago when balloon catheter technology in $\frac{\text{PDT}}{\text{ballon}}$ was not as advanced as it is today. New endovascular light ballon catheters, however, can remove most of the blood. . .
- SUMM [0020] The use of wavelengths of light lower than 600 nm offers significant advantages in PDT because such wavelengths have penetration characteristics that deliver the PDT effect to the target sites (media and adventicia layers of the vessel) and not to myocardial tissue. Thus, effective therapy can be afforded at the target site, while deeper tissues are shielded from a PDT response by blood absorption within these tissues. Previously reported cardiovascular experiments performed to date on tetrapyrrolic molecules have been done. . .
- SUMM . . . lasers are available. At other wavelengths (besides blue)<600 nm-only dye lasers exist to supply enough light power to undertake a PDT treatment. These are particularly useful at 580 nm. Blue lasers are available, and even though most of the photosensitizers that.

SUMM . . . to produce a gallium tetrapyrrolic complex, unexpectedly markedly enhances the uptake and biological efficacy of the compounds as photosensitizers for PDT of cardiovascular diseases when compared to the corresponding tetrapyrrolic compounds having other metal types coordinated to their central cavity. Additionally,. . SUMM [0027] The invention also provides new methods of treating cardiovascular diseases with PDT utilizing light at shorter wavelengths with the new metallated porphyrins of the invention, thus minimizing damage to the myocardial or. . . [0028] The invention further provides new photosensitizers that may be SUMM used in short wavelength applications in photodynamic therapy to treat diseases other than cardiovascular diseases. SUMM . . . the present invention, in one aspect, provides phototherapeutic compositions of metallotetrapyrrolic compounds of formula I which may be used in photodynamic therapy or in a medicament for ##STR1## treatment of diseases such as cardiovascular diseases: . . . of the invention, provided are phototherapeutic compositions of SUMM metallo-tetrapyrrolic compounds of formula II that may be useful as photosensitizers in photodynamic therapy or in a medicament for treatment of diseases such as cardiovascular diseases: . . . accordance with the present invention, provided are SUMM phototherapeutic compositions of metallo-tetrapyrrolic compounds of formula III which may be useful in photodynamic therapy or in a medicament for treatment of diseases such as cardiovascular diseases: ##STR6## . . . accordance with the present invention, provided are SUMM phototherapeutic compositions of metallo-tetrapyrrolic compounds of formula IV which may be used in photodynamic therapy or in a medicament for treatment of diseases such as cardiovascular diseases: ##STR8## . . . synthetic routes to novel tetrapyrrolic molecules of interest SUMM in treating diseases of the cardiovascular system and other diseases applicable to PDT. Such derivatives are of particular interest because all display absorption maximas at wavelengths at or near 400 nm, 532 nm. . . . hydroxylated residue is present. The new porphyrins themselves SUMM may be photodynamically active as metal free analogs and therefore useful as PDT agents. However, metallated derivatives of these compounds are of particular interest in treatment of cardiovascular disease and normal or abnormal. . . SUMM . . . hydroxylated residue is present. The new porphyrins themselves may be photodynamically active as metal free analogs and therefore useful as PDT agents. However, metallated derivatives of these compounds are of particular interest in treatment of cardiovascular disease and normal or abnormal. . . SUMM [0194] 12 week old female albino Hartley guinea pigs (Simonsen:Sim HA) (n=3) were used to assess the effects of $\underline{photodynamic}$ therapy with the gallium tetrapyrroles in gel vehicle applied to the skin. Gallium tetrapyrroles in gel vehicle were administered at 0.1. SUMM [0196] 12 week old male Sprague Dawley rats (Harlan) (n=11) were used to assess the effects of photodynamic therapy with gallium tetrapyrroles (121, 15, 66) in gel vehicle applied to the skin.

Gallium tetrapyrroles in gel vehicle were administered. . .
. . pigmentation, urticaria, allegenic reactions, chronic

hair or hair follicles, disorders of skin pigmentation, acne,

dermatitis, cutaneous vasculitis, erythema multiforme and

cutaneous infections, skin tumors, seborrheic

proliferative dermatitis, chronic ulcerative dermatitis, disorders of

nodosum.

- SUMM . . . and examined by light microscopy to histologically assess the cell population density in the medial and adventitial layers of the $\frac{\text{PDT}}{\text{-treated vessel wall. Tables 3, 4, 5 and 6 contain results}}$ expressed as the % maximum accellularity (depletion of cell population.
- SUMM . . . G. D., Crocker, I. R., Scott, N. A. King, S. B., Wilcox, J. N., Circulation, 96, 1944-1952, 1997). If vascular \underline{PDT} is to be proposed as a therapy to prevent restenosis in humans due to angioplasty or stenting, then it must. . .
- SUMM . . . irradiance) arteries. In another set of experiments, animals also received balloon injuries in the coronary arteries at the time of PDT treatment. Angioplasty injuries in 2 coronary arteries were performed. Vital signs and cardiovascular parameters such as ECG, HR, BP, were. . .
- SUMM [0221] For acute experiments done in uninjured arteries, 3-5 days after the \underline{PDT} experiments, animals were sacrificed and serial sections of all relevant arteries (iliacs, & coronaries) were harvested in 10% formalin and processed for histological assessment. Results of \underline{PDT} at this timepoint give us an insight into the selective cellular effects of \underline{PDT} on VSMC and myofibroblasts which are known to be maximally proliferating and migrating at this same time in response to. . .
- SUMM [0222] For longer term efficacy experiments (14 days after the \underline{PDT} experiments) animals were sacrificed and serial sections of all relevant arteries (coronaries only) were harvested in 10% formalin and processed. . . neointimal formation. The magnitude of the inhibition was greater than any other photosensitizer drug currently used by other groups in \underline{PDT} (clinically or pre-clinically), and was on the order of that only previously seen with radiation in this model. Inhibition data. . .
- SUMM . . . no observed normal skin response at the drug doses used. It has also been noted that significant acellularity occurs following $\frac{\text{PDT}}{\text{azaporphyrins and gallium porphyrins at longer treatment times post injection (16, 24. . .$
- SUMM . . . vascular brachytherapy and to our knowledge are dramatically better than any other photosensitizers described to date in vascular studies with PDT.
- SUMM . . . vascular brachytherapy and to our knowledge are dramatically better than any other photosensitizer described to date in vascular studies with PDT.
- SUMM . . . for example, arc lamps, LEDs or lasers at a certain frequency in the visible spectrum or near infrared for typical PDT treatments. In particular, wavelengths between 400 nm and 900 nm, corresponding to laser diode activation, may also be used. Additionally.
- CLM What is claimed is:
 - . . pigmentation, urticaria, allegenic reactions, chronic proliferative dermatitis, chronic ulcerative dermatitis, disorders of hair or hair follicles, disorders of skin pigmentation, acne, cutaneous infections, skin tumors, seborrheic dermatitis, cutaneous vasculitis, erythema multiforme or nodosum.

L16 ANSWER 6 OF 9 USPATFULL on STN

ACCESSION NUMBER: 2003:147005 USPATFULL

TITLE: Substituted porphyrin and azaporphyrin derivatives and

their use in photodynamic therapy, radioimaging and MRI diagnosis

INVENTOR(S): Robinson, Byron C., Santa Barbara, CA, UNITED STATES NUMBER KIND DATE _____ ___ US 20030100752 A1 20030529 US 6906050 B2 20050614 US 2002-159580 A1 20020531 (10) PATENT INFORMATION: APPLICATION INFO.: NUMBER DATE _____ PRIORITY INFORMATION: US 2001-295343P 20010531 (60) DOCUMENT TYPE: Utility
FILE SEGMENT: APPLICATION LEGAL REPRESENTATIVE: Finnegan, Henderson, Farabow,, Garrett & Dunner, LLP, 1300 I Street, N.W., Washington, DC, 20005-3315 120 NUMBER OF CLAIMS: EXEMPLARY CLAIM: 1 LINE COUNT: 4498 CAS INDEXING IS AVAILABLE FOR THIS PATENT. Substituted porphyrin and azaporphyrin derivatives and their use in photodynamic therapy, radioimaging and MRI diagnosis . azaporphyrin deviations with various substitutents in the 12-AB and 17-positions of the porphyrin skeleton as pharmaceutical agents for use in photodynamic therapy, MRI diagnosis, and radiodiagnostics. . . . derivatives with various substituents in the 13- and SUMM 17-positions of the porphyrin skeleton suitable as pharmaceutical agents for use in photodynamic therapy, MRI diagnosis, and radiodiagnostics. The invention is also directed to pharmaceutical agents that contain these compounds, as well as a. . . [0002] Photodynamic therapy ("PDT") is a SUMM new modality for the treatment of malignancies, diseased tissue, hyperproliferating tissues, pathogens or unwanted normal tissues. PDT involves a localized or systemic administration of a photosensitizing compound followed by exposure of target tissue to photoactivating light. The. . . [0004] Porphyrins and azaporphyrins and their metallated derivatives SUMM belong to a family of substances that are suitable for PDT. These compounds accumulate in target tissues and absorb light in a range in which living tissue is still fairly permeable,. . . have been developed largely for use in oncological applications, but have also been examined in other disease areas in the PDT field in humans. (WO 92/06097; WO 97/20846; EP 0 8116 $\overline{26}$; U.S. Pat. Nos. 5,633,275, 5,654,423, 5,675,001, 5,703,230, and 5,705,622). Such photosensitizers include Photofrin (U.S. Pat. No. 4,882,234), 5-aminolevulinic acid (protoporphyrin IX precursor), SnET2, <u>Visudyne</u>® (Benzoporphyrin derivative), Antrin®, Optrin® (Lutetium texaphyrin) and mono-aspartyl chlorin e6 (MACE). All of these compounds were designed specifically for the. . . SUMM . . . than 600 nm in the cardiovascular field. This may have been true several years ago when balloon catheter technology in PDT was not as advanced as it is today. New endovascular light $\overline{\text{bal}}$ lon catheters, however, can remove most of the blood. $\ . \ \ .$ SUMM [0008] The use of wavelengths of light lower than 600 nm offers significant advantages in PDT because such wavelengths have penetration characteristics that deliver the PDT effect to the

target sites (media and adventicia layers of the vessel) and not to myocardial tissue. Thus, effective therapy can be afforded at the target

site, while deeper tissues are shielded from a PDT response by

- blood absorption within these tissues. Previously reported cardiovascular experiments performed to date on tetrapyrrolic molecules have been done. . .
- SUMM . . . However, the compounds so far described are far from being able to satisfactorily meet the desired requirements to be effective PDT, MRI and radiodiagnostic imaging agents.
- SUMM . . . providing metalloporphyrin amide linkages. However, all of these approaches using deuteroporphyrins are suboptimal with respect to design of short wavelength PDT photosensitizers for use as MRI or radiodiagnostic agents for reasons detailed below.
- SUMM [0015] Sakata's porphyrin-based $\underline{PDT}/MRI/radiodiagnostic$ compounds are based on a naturally occurring asymmetrical porphyrin ring system shown in FIG. 1.
- SUMM . . . absorptions at about 532 and 575 nm with molar extinction coefficients of between 15,000-20,000 M.sup.-1 cm.sup.-1. In the field of photodynamic therapy, the depth of light penetration into tissues is a function of the wavelength of the exciting light. The theoretical efficacy. . .
- SUMM . . . the properties and uses of the compounds clinically for not only MRI and radiodiagnostic imaging, but also for treatment using photodynamic therapy.
- SUMM . . . found novel metal-free or metallated functionalized phototherapeutic agents that may be used for imaging (MRI or radiodiagnostic) before or after photodynamic therapy . These novel phototherapeutic agents are based on tetrapyrrolic ring systems such as the porphyrins and azaporphyrins that can be covalently linked by stable linkages to metal complexing agents. These new photosensitizers are useful in short wavelength applications in photodynamic therapy.
- SUMM . . . in one aspect provides phototherapeutic compositions of metallotetrapyrrolic compounds of formula I which may be used as MRI, radiodiagnostic and PDT agents: ##STR2##
- SUMM . . . provided are phototherapeutic, MRI and radiodiagnostic compositions of metallo-tetrapyrrolic compounds of formula II that may be used as photosensitizers in photodynamic therapy:
 ##STR8##
- SUMM . . . present invention, provided are phototherapeutic compositions of metallo-tetrapyrrolic compounds of formula III that may be used as MRI, radiodiagnostic, or PDT agents: ##STR12##
- SUMM . . . the invention, $\overline{\text{prov}}$ ided are phototherapeutic compositions of metallo-tetrapyrrolic compounds of formula IV that may be used as MRI, radiodiagnostic, or PDT agents: ##STR16##
- SUMM . . . the metal co-ordination compound. The new porphyrins themselves may be photodynamically active as metal free analogs and therefore useful as \underline{PDT} agents. In addition, metallated derivatives of these compounds are also of particular interest for treatment and diagnosis of disorders of. . .
- SUMM [0182] In accordance with the invention, the porphyrin or azaporphyrin linked MCR compounds can then be modified to produce \underline{PDT}/MRI radiodiagnostic compounds. If the compounds are to be used for NMR diagnosis, paramagnetic metal ions must be present in the. . .
- SUMM [0183] For the use of the agents according to the invention for $\frac{\text{photodynamic therapy}}{\text{compound should be metal free, i.e, }M=2\text{H, or should have coordinated}}$ photoactive metals, preferred examples of. . .
- SUMM . . . and are generally dosed in amounts of 0.01 μ mol to 2 mmol/kg of body weight, both for their use in \underline{PDT} and for therapy monitoring using MRI diagnosis. They are intended for enteral and parenteral administration or are administered with the. . .
- SUMM [0194] The agents according to the invention are especially suitable for

SUMM

SUMM

PDT and as MRI contrast media. After administration, they can enhance the informational value of the image that is obtained from. .

. . . in addition to therapeutics. Additionally, as more disease indications are realized, shorter wavelength light may be equally important in other PDT applications that only require short wavelength excitation to effect a therapy. Such applications may be, for example, in hollow organ disease (for example lung cancers, barrets esophagus), or in diseases of the skin (for example psoriasis, actinic keratosis, acne vulgaris). The invention disclosed herein describes the synthesis of metallated photosensitizers having ring systems that have shown excellent efficacy in. . . clearance characteristics and low toxicity. (See co-pending application filed on May 31, 2001 entitled "Metallotetrapyrrolic Photosensitizing Agents For Use In Photodynamic Therapy," inventors Byron C. Robinson, Ian M. Leitch, Stephanie Greene, and Steve Rychnovsky,

Attorney Docket No. 07328-0015.)

effective $\underline{photodynamic}$ $\underline{\underline{therapy}}$ treatment but also as MRI and radiodiagnostic diagostic agents. Such compounds may be used to diagnose, locate or treat cardiovascular. . .

[0250] The compounds of the invention are intended for use not only for

L16 ANSWER 7 OF 9 USPATFULL on STN

2003:44768 USPATFULL ACCESSION NUMBER:

TITLE: Methods and compositions for the treatment of macular

and retinal degenerations

Travis, Gabriel H., Los Angeles, CA, UNITED STATES INVENTOR(S): PATENT ASSIGNEE(S): Board of Regents, The University of Texas System (U.S.

corporation)

NUMBER KIND DATE -----PATENT INFORMATION: US 20030032078 A1 20030213 US 2001-885303 A1 20010619 <--A1 20010619 (9) APPLICATION INFO.:

> NUMBER DATE _____

PRIORITY INFORMATION: US 2001-263837P 20010123 (60)

PRIORITY INFORMATION

DOCUMENT TYPE: Utility

APPLICATION

Chic

LEGAL REPRESENTATIVE: Gina N. Shishima, Fulbright & Jaworski L.L.P., Suite

2400, 600 Congress Avenue, Austin, TX, 78701

NUMBER OF CLAIMS: 53 1 EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 7 Drawing Page(s)

LINE COUNT: 7372

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

. . . for the prevention and treatment of macular and retinal degeneration. Early detection of macular degeneration is also becoming increasingly important. Photodynamic therapy, a surgical treatment for some cases of macular degeneration, is only beneficial before extensive vision loss has occurred (Bressler, et. .

SUMM . . . other length of time wherein repeating the therapy is necessary. In some embodiments, surgery such as laser photocoagulation therapy or photodynamic therapy may be performed on the subject or an anti-angiogenic factor may be administered to the subject. An "effective amount" refers. . .

. . . phosphodiester backbone moiety used for improved nuclease DETD

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resistance, cellular uptake and regulating RNA expression; U.S. Pat. No.
       5,858,988 which describes hydrophobic carrier agent attached
       to the 2'-O position of oligonucleotides to enhanced their membrane
       permeability and stability; U.S. Pat. No. 5,214,136.
DETD
       [0408] Treatments developed that reduce the risk of vision loss in
       selected patients with "wet" macular degeneration include
       photocoagulation and photodynamic therapy. These
       therapies may be used in conjunction with a therapeutic agent which has
       been through screening using a dehydrogenase.
DETD
       [0410] Photodynamic therapy, allows for the
       treatment of patients with neovascular macular degeneration having
       vessels extending under the center of the retina. Photodynamic
       therapy uses the drug verteporfin, and has recently
       been shown to reduce the risk of moderate and severe vision loss
       (Treatment of Age-Related Macular Degeneration with Photodynamic
       Therapy (TAP) Study Group, 1999, 2000) In photodynamic
       therapy, a photoactivator, verteporfin, is injected
       into a patients vein where it then travels to the eye and becomes
       concentrated within the neovascular lesion. Then a laser is applied over
       the entire neovascular lesion to activate the drug. The photoactivated
       verteporfin selectively destroys lesions by creating reactive
       intermediates of oxygen such as superoxide and hydroxide radicals
       without damaging viable retinal tissue. . . et al., 2000).
       Retreatment as often as every three months are needed to prevent
       significant growth. The laser used in photodynamic
       therapy is not a "heat producing" laser as used in
       photocoagulation. Generally, this therapy works for blood vessels that
       are not. . . fluid in growths wherein the neovascularization is less
       than about 50% (www.macular-degeneration.org/porphyrin/porphyrin.html).
       Clinical trials have shown that photo-dynamic therapy with
       verteporfin could reduce the risk of moderate and severe vision
       loss from 61% to 33% at one year and from 69%. . .
DETD
       . . of 11cRDH in RPE cells (Law et al., 1989; Gamble et al., 1999).
       This drug is used clinically to treat <a href="mailto:acne">acne</a> because of
       unrelated effects on sebaceous glands. Night blindness is a common side
       effect of isotretinoin due to impaired regeneration. .
       . . . optimal dose of isotretinoin to block formation of A2E in
DETD
       abcr-/- mice can be determined. The recommended dose for treating
       acne is 0.5 to 2.0~mg/kg/day. The observation of occasional
       night blindness in humans suggests significant impairment of rhodopsin
       regeneration at. . . in RPE tissue should be achieved at doses
       similar to, or possibly below, human therapeutic doses for the treatment
       of acne. The effects of isotretinoin on atRAL in retinas from
       light-adapted mice would preferably be determined at doses that bracket
       the. . .
DETD
       [0501] Bressler N M, Gills J P. Age related macular degeneration. New
       hope for a common problem comes from photodynamic
       therapy. BMJ Dec. 9, 2000;321(7274):1425-1427.
DETD
       [0584] Hasan T, Schmidt-Erfurth U. Mechanisms of action of
       photodynamic therapy with verteporfin for
       the treatment of age-related macular degeneration. Surv Ophthalmol (in
       [0800] Treatment of Age-Related Macular Degeneration with
DETD
       Photodynamic Therapy (TAP) Study Group.
       Verteporfin (VisudyneJ) therapy of subfoveal choroidal
       neovascularization in age-related macular degeneration. One year results
       of two randomized clinical trials: TAP report. . .
DETD
       [0801] Treatment of Age-Related Macular Degeneration With
       Photodynamic Therapy (TAP) Study Group.
       Photodynamic therapy of subfoveal choroidal
```

CLM

neovascularization in age-related macular degeneration with verteporfin: two-year results of 2 randomized clinical trials: TAP report No 2. Arch Ophthalmol (in press). What is claimed is:

52. The method of claim 39, further comprising performing photodynamic therapy on the subject.

L16 ANSWER 8 OF 9 USPATFULL on STN

2001:124629 USPATFULL ACCESSION NUMBER:

Photoactivation of endogenous porphyrins for treatment TITLE:

of psoriasis

INVENTOR(S): Lui, Harvey, Vancouver, Canada

Macaulay, Calum, Vancouver, Canada Zeng, Haishan, Delta, Canada

McLean, David I., Vancouver, Canada

Bissonnette, Robert, Vancouver, Canada

PATENT ASSIGNEE(S): The University of British Columbia, Vancouver, Canada

(non-U.S. corporation)

NUMBER KIND DATE _____

PATENT INFORMATION: US 6269818 B1 20010807 APPLICATION INFO.: US 1998-84865 19980526 DOCUMENT TYPE: Utility <--

19980526 (9)

DOCUMENT TYPE: FILE SEGMENT: GRANTED

PRIMARY EXAMINER: Peffley, Michael ASSISTANT EXAMINER: Kearney, R.

LEGAL REPRESENTATIVE: Townsend and Townsend and Crew LLP

NUMBER OF CLAIMS: 19 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 21 Drawing Figure(s); 11 Drawing Page(s)

LINE COUNT: 1053

Autofluorescence photographic images have been used to evaluate treatment responses in acne (Lucchina et al 1996, Martin R. J.

et al 1973). Analysis and comparison of emission spectra has also been studied. .

SUMM

. . under Wood's lamp illumination was reported as early as 1927(Bommer, 1927), and has been linked to the presence in acne of porphyrins generated by Propionibacterium acnes (Cornelius, 1967;

McGinley, 1980; Lee et al 1978; Konig et al, 1992; Johnson, 1987;. .

Macrospectrophotometry may be used to detect skin porphyrin in patients receiving exogenous porphyrins, or porphyrin precursors, for

photodynamic therapy, and to follow the time course accumulation of porphyrins in photodynamic therapy

(Lui, 1996; Rhodes, 1997; Stringer, 1996). The intensity of the

fluorescence emission peaks has been shown to correlate with the. . . SUMM . . (Arakane et al., 1996). The toxicity generated by light

activation of pharmacologically elevated levels of porphyrins is the

basis for photodynamic therapy which may be used to treat a variety of conditions, including cancer (see U.S. Pat. Nos.

5,211,938; 5,234,940; 5,079,262; all. . .

. . . Goerz et al., 1995, report that skin does not normally contain SUMM

sufficient levels of porphyrins to allow one to perform

photodynamic therapy, and consequently

photodynamic therapy requires exogenous addition of photosensitizer.

DETD TABLE II

Clinical diagnosis of patients studied

Diagnosis

```
Psoriasis
                         11
       Contact dermatitis
      Atopic dermatitis
        <u>Seborrheic</u> <u>dermatitis</u> 2
                                10
                              12
      Actinic keratosis
                              18
      Port wine stain
                             3
                             3
      Porokeratosis
       Discoid lupus erythematosus 2
       Rosacea
       Sebaceous hyperplasia. .
DETD
      Boehncke, W. H., Sterry, W. & Kaufmann, R. (1994). Treatment of
      psoriasis by topical photodynamic therapy with
      polychromatic light [letter]. Lancet, 343:801.
DETD
      Goff, B. A., Bachor, R., Kollias, N. & Hasan, T. (1992). Effects of
      photodynamic therapy with topical application of
       5-aminolevulinic acid on normal skin of hairless guinea pigs. Journal of
       Photochemistry & Photobiology. B -. . .
       Gudgin Dickson, E. F. & Pottier, R. H. (1995). On the role of
       protoporphyrin IX photoproducts in photodynamic
       therapy [news]. Journal of Photochemistry & Photobiology, B -
      Biology, 29:91-3.
      . . . In-vivo fluorescence detection and imaging of
DETD
      porphyrin-producing bacteria in the human skin and in the oral cavity
      for diagnosis of acne vulgaris, caries, and squamous cell
       carcinoma. SPIE 2135:129.
DETD
      Lucchina, L. et al. (1996). Fluorescence photography n the evaluation of
      acne. J. Am Acad Dermatol 35:58-63.
       . . H., Zeng, H., McLean, D. I., MacAulay, C. E. & Palcic, B.
DETD
       (1996). In vivo fluorescence spectroscopy monitoring of BPD
       verteporfin concentration changes in skin tissue during
       photodynamic skin cancern. Journal of Dermatological Science, 12:87.
DETD
      Nelson, L. S. et al. (1985). Tropical 5-aminolevulinic acid (ALA) for
       the photodynamic therapy of psoriasis and actinic
       keratoses. Am. Soc. For Laser Medicine and Surgery Abstracts, p. 43,
      Abstract 202.
       . . (1996). The accumulation of Protoporphyrin Ix in Plaque
DETD
      Psoriasis After Topical Application of 5-Aminolevulinic Acid Indicated a
       Potential For superficial Photodynamic Therapy.
      Journal of Investigative Dermatology, 107:76-81.
      Szeimies, R., Calzavara-Pinton, P., Karrer, S., Ortel, B. and
      Landthaler, M. (1996). Topical photodynamic therapy
       in dermatology. J. of photochemistry and photobiology 36:213-219.
DETD
      Tan, W. C., Krasner, N., O'Toole, P. and Lombard, M. (1997). Enhancement
       of photodynamic therapy in gastric cancer cells by
       removal of iron. Gut 41:14-18.
L16 ANSWER 9 OF 9 USPATFULL on STN
                       93:69868 USPATFULL
ACCESSION NUMBER:
TITLE:
                       Compositions for photodynamic therapy
                       Liu, Daniel, Vancouver, Canada
INVENTOR(S):
                       Jiang, Frank, Vancouver, Canada
                       Hobbs, John, Vancouver, Canada
                       Quadra Logic Technologies Inc., Vancouver, Canada
PATENT ASSIGNEE(S):
                       (non-U.S. corporation)
                          NUMBER KIND DATE
                       ______
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Number of Patients

PATENT INFORMATION: US 5238940 19930824 <--

US 1991-768810 APPLICATION INFO.: 19910930 (7)

RELATED APPLN. INFO.: Continuation-in-part of Ser. No. US 1990-498042, filed

on 22 Mar 1990, now patented, Pat. No. US 5053423

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

Raymond, Richard L. PRIMARY EXAMINER: LEGAL REPRESENTATIVE: Morrison & Foerster

NUMBER OF CLAIMS: 8 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 24 Drawing Figure(s); 16 Drawing Page(s)

LINE COUNT: 1191

CAS INDEXING IS AVAILABLE FOR THIS PATENT. ΤТ Compositions for photodynamic therapy

SUMM The present invention relates to methods to prepare pharmaceutical compositions useful in photodynamic therapy. More specifically, the invention concerns conjugates of porphyrin-type photosensitizers with hydrophilic polymers as active ingredients in compositions which can be. . .

SUMM The products of the invention method are pharmaceutical compositions useful in photodynamic therapy or related methodologies, which compositions contain as an active ingredient a conjugate of a porphyrin-type photosensitizer with a water soluble,.

DETD An additional group of compounds which has been found extremely useful in photodynamic therapy and related methodologies is the green porphyrin (Gp) group having the basic structure outlined in FIG. 1. These compounds are. . .

DETD . . . solid tumors, dissolution of plaques in blood vessels (see, e.g., U.S. Pat. No. 4,512,762); treatment of topical conditions such as acne, athlete's foot, warts, papilloma, and psoriasis and treatment of biological products (such as blood for transfusion) for infectious agents, since. . .

TТ 9002-89-5D, modified, conjugates with porphyrin derivs. Photofrin II, conjugates with modified polyvinyl alc. 129497-78-5D, conjugates with modified polyvinyl alc. (as photosensitizer for photodynamic therapy)